

AMENDMENTS TO THE CLAIMS

Please cancel claims 1 to 23, without prejudice.

Please amend the claims as follows.

Please add new claims 45 to 79.

Claims 1 to 23 (canceled)

24. (currently amended) A method for making an aqueous solution of metronidazole greater than 0.75% w/w comprising combining metronidazole, beta-cyclodextrin (BCD), and niacin or niacinamide in water.

25. (original) The method of claim 24 wherein the metronidazole is added to the water after the BCD and the niacin or niacinamide are dissolved in the water.

26. (original) The method of claim 24 which further comprises, after the combination of metronidazole, BCD, and the niacin or niacinamide, adding a gelling agent.

27. (original) The method of claim 24 wherein niacinamide but not niacin is combined.

28. (original) The method of claim 24 wherein niacin but not niacinamide is combined.

29. (original) An aqueous solution that is made by the method of claim 24.
30. (original) An aqueous solution that is made by the method of claim 26.
31. (original) An aqueous solution that is made by the method of claim 27.
32. (original) An aqueous solution that is made by the method of claim 28.
33. (original) A method for the treatment of a dermatologic or mucosal disorder comprising topically applying an effective amount of aqueous solution of metronidazole having a concentration higher than 0.75% w/w to the site of the disorder and permitting the metronidazole to treat the disorder, wherein the solution comprises beta-cyclodextrin (BCD) and niacin or niacinamide.
34. (original) The method of claim 33 wherein the concentration of metronidazole is about 1% or higher.
35. (original) The method of claim 34 wherein the application is once daily.
36. (original) The method of claim 33 wherein the disorder is rosacea.

37. (original) The method of claim 33 wherein the solution comprises niacin and is substantially free of niacinamide.

38. (original) The method of claim 33 wherein the solution comprises niacinamide and is substantially free of niacin.

39. (original) The method of claim 33 wherein the aqueous solution is a gel.

40. (currently amended) A kit for the topical treatment of dermatologic or mucosal disorders comprising a container and an aqueous solution of metronidazole, beta-cyclodextrin, and niacin or niacinamide within said container, wherein the concentration of metronidazole in said solution is higher than 0.75% w/w.

41. (currently amended) The kit of claim 40 wherein the concentration of metronidazole is about 1% w/w or higher.

42. (original) The kit of claim 40 wherein the aqueous solution is a gel.

43. (original) The kit of claim 40 wherein the solution contains niacinamide and is substantially free of niacin.

44. (original) The kit of claim 40 wherein the solution contains niacin and is substantially free of niacinamide.

45. (new) An aqueous solution that is physically stable for at least one week at 5°C comprising metronidazole, betacyclodextrin, and a solubility enhancing agent which is niacin or niacinamide.

46. (new) The aqueous solution of claim 45 wherein the solubility enhancing agent is niacinamide.

47. (new) The aqueous solution of claim 46 wherein the solubility enhancing agent is niacin.

48. (new) The aqueous solution of claim 45 wherein the concentration of metronidazole in the solution is higher than 0.75% w/w.

49. (new) The aqueous solution of claim 48 wherein the concentration of metronidazole is about 1.0% or higher.

50. (new) The aqueous solution of claim 48 wherein the solubility enhancing agent is niacinamide.

51. (new) The aqueous solution of claim 48 wherein the solubility enhancing agent is niacin.

52. (new) The aqueous solution of claim 45 wherein in the solution, the concentration of metronidazole is about 1.0% w/w, the concentration of betacyclodextrin is about 1.0% w/w or less, and the concentration of niacinamide or niacin is about 0.5% w/w or more.

53. (new) The aqueous solution of claim 52 which comprises niacinamide and does not comprise niacin.

54. (new) The aqueous solution of claim 52 wherein the concentration of niacinamide or niacin is about 1.0% w/w or higher.

55. (new) The aqueous solution of claim 54 which comprises niacinamide and does not comprise niacin.

56. (new) The aqueous solution of claim 46 wherein the concentration of niacinamide in the solution is below that which, in the absence of betacyclodextrin, increases the water solubility of metronidazole to the concentration of metronidazole present in the solution.

57. (new) The aqueous solution of claim 47 wherein the concentration of niacin in the solution is below that which, in the absence of betacyclodextrin, increases the water solubility of metronidazole to the concentration of metronidazole present in the solution.

58. (new) The aqueous solution of claim 45 wherein the concentration of betacyclodextrin in the solution is below that which, in the absence of niacinamide or niacin, increases the water solubility of metronidazole to the concentration of metronidazole present in the solution.

59. (new) The aqueous solution of claim 1 which is a gel.

60. (new) The aqueous solution of claim 48 which is a gel.

61. (new) The aqueous solution of claim 50 which is a gel.

62. (new) The aqueous solution of claim 51 which is a gel.

63. (new) A method for increasing the solubility of metronidazole in aqueous solution comprising combining metronidazole, betacyclodextrin, and niacinamide or niacin in an aqueous fluid.

64. (new) The method of claim 63 wherein the aqueous solution is a gel.

65. (new) The method of claim 63 which comprises combining niacinamide in the fluid.

66. (new) The method of claim 63 which comprises combining niacin in the fluid.

67. (new) The method of claim 63 wherein the solubility of metronidazole is increased to 0.75% w/w or more.

68. (new) The method of claim 67 wherein the solubility of metronidazole is increased to about 1.0% w/w or more.

69. (new) The method of claim 63 wherein the betacyclodextrin and the niacin and niacinamide are dissolved in the aqueous fluid before the metronidazole is combined in the fluid.

70. (new) The method of claim 63 wherein a gelling agent is added to the fluid after the metronidazole, betacyclodextrin, and the niacin or niacinamide are combined in the fluid.

71. (new) The method of claim 63 wherein the amount of niacinamide or niacin that is combined in the fluid is below that which, in the absence of betacyclodextrin, increases the water solubility of metronidazole to the concentration of metronidazole that is soluble in the solution.

72. (new) The method of claim 63 wherein the amount of betacyclodextrin in the solution that is combined in the fluid is below that which, in the absence of niacinamide or niacin, increases the water solubility of metronidazole to the concentration of metronidazole that is soluble in the solution.

73. (new) The method of claim 63 wherein the amount of niacinamide or niacin that is combined in the fluid is below that which, in the absence of betacyclodextrin, increases the water solubility of metronidazole to the concentration of metronidazole that is soluble in the solution, and the amount of betacyclodextrin in the solution that is combined in the fluid is below that which, in the absence of niacinamide or niacin, increases the water solubility of metronidazole to the concentration of metronidazole that is soluble in the solution.

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74. (new) A method for increasing the enhancing effect of betacyclodextrin on the solubility of metronidazole in aqueous fluid comprising combining niacin or niacinamide with the betacyclodextrin in the aqueous fluid.

75. (new) The method of claim 74 wherein the niacin or niacinamide is combined in the fluid with the betacyclodextrin and then metronidazole is added to the fluid.

76. (new) The method of claim 74 wherein niacin is combined with the betacyclodextrin in the aqueous fluid.

77. (new) The method of claim 74 wherein niacinamide is combined with the betacyclodextrin in the aqueous fluid.

78. (new) The method of claim 74 wherein the concentration of betacyclodextrin in the fluid is about 1.0% or less and the concentration of niacin or niacinamide is about 0.5% w/w or more.

79. (new) The method of claim 78 wherein the concentration of niacin or niacinamide is about 1.0% or more.

80. (new) The method of claim 24 wherein the betacyclodextrin is crystalline betacyclodextrin.

81. (new) An aqueous solution that is made by the method of claim 80.
